09/7236/6

From the INTERNATIONAL SEARCHING AUTHORITY	PCT
Attn. Noonan, Kevin E. 300 South Wacker Drive, Suite 3200 CHICAGO. IL 60606	NOTIFICATION OF TRANSMITTAL OF OR THE INTERNATIONAL SEARCH REPORT OR THE DECLARATION MAY 2 2 2001 File JOS (PCT Rule 44.1) ATE: 7/19/01 8/18/01 Date of mailing (day/month/year) 18/05/2001
Applicant's or agent's file reference	10/03/2001
MBHB00-1214	FOR FURTHER ACTION See paragraphs 1 and 4 below
International application No. PCT/US 00/32400	International filing date (day/month/year) 28/11/2000
AMGEN, INC. et al.	
The applicant is hereby notified that the International Search Filing of amendments and statement under Article 19: The applicant is entitled, if he so wishes, to amend the claim When? The time limit for filing such amendments is normal	ns of the International Application (see Rule 46):
International Search Report; however, for more de Where? Directly to the International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland Fascimile No.: (41-22) 740.14.35	tails, see the notes on the accompanying sheet.
For more detailed instructions, see the notes on the acco	mpanying sheet.
2. The applicant is hereby notified that no International Search Article 17(2)(a) to that effect is transmitted herewith.	n Report will be established and that the declaration under
3. With regard to the protest against payment of (an) addition	nal fee(s) under Rule 40.2, the applicant is notified that:
the protest together with the decision thereon has been applicant's request to forward the texts of both the protest.	n transmitted to the International Bureau together with the lest and the decision thereon to the designated Offices.
no decision has been made yet on the protest; the app	licant will be notified as soon as a decision is made.
4. Further action(s): The applicant is reminded of the following:	
Shortly after 18 months from the priority date, the international ap If the applicant wishes to avoid or postpone publication, a notice priority claim, must reach the International Bureau as provided i completion of the technical preparations for international publica	of withdrawal of the international application, or of the in Rules 90 <i>bis</i> .1 and 90 <i>bis</i> .3, respectively, before the
Within 19 months from the priority date, a demand for international wishes to postpone the entry into the national phase until 30 mo	al preliminary examination must be filed if the applicant nths from the priority date (in some Offices even later).
Within 20 months from the priority date, the applicant must perfor before all designated Offices which have not been elected in the priority date or could not be elected because they are not bound	e demand or in a later election within 19 months from the
Name and mailing address of the International Searching Authority	Authorized officer ,
European Patent Office, P.B. 5818 Patentlaan 2 NL-2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016	Renate Jordan

NOTES TO FORM PCT/ISA/220

These Notes are intended to give the basic instructions concerning the filing of amendments under article 19. The Notes are based on the requirements of the Patent Cooperation Treaty, the Regulations and the Administrative Instructions under that Treaty. In case of discrepancy between these Notes and those requirements, the latter are applicable. For more detailed information, see also the PCT Applicant's Guide, a publication of WIPO.

In these Notes, "Article", "Rule", and "Section" refer to the provisions of the PCT, the PCT Regulations and the PCT Administrative Instructions respectively.

INSTRUCTIONS CONCERNING AMENDMENTS UNDER ARTICLE 19

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The applicant has, after having received the international search report, one opportunity to amend the claims of the international application. It should however be emphasized that, since all parts of the international application (claims, description and drawings) may be amended during the international preliminary examination procedure, there is usually no need to file amendments of the claims under Article 19 except where, e.g. the applicant wants the latter to be published for the purposes of provisional protection or has another reason for amending the claims before international pbulication. Furthermore, it should be emphasized that provisional protection is available in some States only.

What parts of the international application may be amended?

Under Article 19, only the claims may be amended.

During the international phase, the claims may also be amended (or further amended) under Article 34 before the International Preliminary Examining Authority. The description and drawings may only be amended under Article 34 before the International Examining Authority.

Upon entry into the national phase, all parts of the international application may be amended under Article 28 or, where applicable, Article 41.

When?

Within 2 months from the date of transmittal of the international search report or 16 months from the priority date, whichever time limit expires later. It should be noted, however, that the amendments will be considered as having been received on time if they are received by the International Bureau after the expiration of the applicable time limit but before the completion of the technical preparations for international publication (Rule 46.1).

Where not to file the amendments?

The amendments may only be filed with the International Bureau and not with the receiving Office or the International Searching Authority (Rule 46.2).

Where a demand for international preliminary examination has been its filed, see below.

How?

Either by cancelling one or more entire claims, by adding one or more new claims or by amending the text of one or more of the claims as filed.

A replacement sheet must be submitted for each sheet of the claims which, on account of an amendment or amendments, differs from the sheet originally filed.

All the claims appearing on a replacement sheet must be numbered in Arabic numerals. Where a claim is cancelled, no renumbering of the other claims is required. In all cases where claims are renumbered, they must be renumbered consecutively (Administrative Instructions, Section 205(b)).

The amendments must be made in the language in which the international application is to be published.

What documents must/may accompany the amendments?

Letter (Section 205(b)):

The amendments must be submitted with a letter.

The letter will not be published with the international application and the amended claims. It should not be confused with the "Statement under Article 19(1)" (see below, under "Statement under Article 19(1)").

The letter must be in English or French, at the choice of the applicant. However, if the language of the international application is English, the letter must be in English; if the language of the international application is French, the letter must be in French.

NOTES TO FORM PCT/ISA/220 (continued)

The letter must indicate the differences between the claims as filed and the claims as amended. It must, in particular, indicate, in connection with each claim appearing in the international application (it being understood that identical indications concerning several claims may be grouped), whether

- (i) the claim is unchanged;
- (ii) the claim is cancelled:
- (iii) the claim is new;
- (iv) the claim replaces one or more claims as filed;
- (v) the claim is the result of the division of a claim as filed.

The following examples illustrate the manner in which amendments must be explained in the accompanying letter:

- [Where originally there were 48 claims and after amendment of some claims there are 51]:
 "Claims 1 to 29, 31, 32, 34, 35, 37 to 48 replaced by amended claims bearing the same numbers; claims 30, 33 and 36 unchanged; new claims 49 to 51 added."
- [Where originally there were 15 claims and after amendment of all claims there are 11]: "Claims 1 to 15 replaced by amended claims 1 to 11."
- [Where originally there were 14 claims and the amendments consist in cancelling some claims and in adding new claims]:
 "Claims 1 to 6 and 14 unchanged; claims 7 to 13 cancelled; new claims 15, 16 and 17 added." or "Claims 7 to 13 cancelled; new claims 15, 16 and 17 added; all other claims unchanged."
- [Where various kinds of amendments are made]:
 "Claims 1-10 unchanged; claims 11 to 13, 18 and 19 cancelled; claims 14, 15 and 16 replaced by amended claim 14; claim 17 subdivided into amended claims 15, 16 and 17; new claims 20 and 21 added."

"Statement under article 19(1)" (Rule 46.4)

The amendments may be accompanied by a statement explaining the amendments and indicating any impact that such amendments might have on the description and the drawings (which cannot be amended under Article 19(1)).

The statement will be published with the international application and the amended claims.

It must be in the language in which the international appplication is to be published.

It must be brief, not exceeding 500 words if in English or if translated into English.

It should not be confused with and does not replace the letter indicating the differences between the claims as filed and as amended. It must be filed on a separate sheet and must be identified as such by a heading, preferably by using the words "Statement under Article 19(1)."

It may not contain any disparaging comments on the international search report or the relevance of citations contained in that report. Reference to citations, relevant to a given claim, contained in the international search report may be made only in connection with an amendment of that claim.

Consequence if a demand for international preliminary examination has already been filed

If, at the time of filing any amendments under Article 19, a demand for international preliminary examination has already been submitted, the applicant must preferably, at the same time of filing the amendments with the International Bureau, also file a copy of such amendments with the International Preliminary Examining Authority (see Rule 62.2(a), first sentence).

Consequence with regard to translation of the international application for entry into the national phase

The applicant's attention is drawn to the fact that, where upon entry into the national phase, a translation of the claims as amended under Article 19 may have to be furnished to the designated/elected Offices, instead of, or in addition to, the translation of the claims as filed.

For further details on the requirements of each designated/elected Office, see Volume II of the PCT Applicant's Guide.

PATENT COOPERATION TREATY





(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference MBHB00-1214	FOR FURTHER ACTION			ational Search Report applicable, item 5 below.
International application No.	International filing date (day	y/month/year)	(Earliest) Priority D	ate (day/month/year)
PCT/US 00/32400	28/11/200	00	10/	12/1999
Applicant				
AMGEN, INC. et al.		· .		
This International Search Report has been according to Article 18. A copy is being tra	n prepared by this Internation Insmitted to the International	al Searching Autho Bureau.	rity and is transmitte	d to the applicant
This International Search Report consists X It is also accompanied by	of a total of5 a copy of each prior art docu	sheets. ment cited in this re	eport.	
1. Basis of the report			,	
 a. With regard to the language, the language in which it was filed, unl 			of the international	application in the
the international search w Authority (Rule 23.1(b)).	as carried out on the basis of	a translation of the	international applica	ation furnished to this
[]	e sequence listing : nal application in written form	1.	rnational application	, the international search
<u> </u>	rnational application in compi this Authority in written form.			
	this Authority in computer rea			
	sequently furnished written s		s not go beyond the	disclosure in the
the statement that the info furnished	rmation recorded in compute	r readable form is i	dentical to the writter	n sequence listing has been
2. X Certain claims were four	nd unsearchable (See Box I)) .		
3. Unity of invention is lack	king (see Box II).	•		
4. With regard to the title,				
X the text is approved as su	emitted by the applicant.			
the text has been establish	ned by this Authority to read a	as follows:		
5. With regard to the abstract,			,	
the text is approved as sult the text has been establish within one month from the	omitted by the applicant. ned, according to Rule 38.2(b date of mailing of this interna	b), by this Authority attional search repor	as it appears in Box t, submit comments	III. The applicant may, to this Authority.
6. The figure of the drawings to be publi	-	ıre No.		·
as suggested by the applic				None of the figures.
because the applicant falle because this figure better	ed to suggest a figure. characterizes the invention.			

International A tion No PCT/US 00/32400

A. CLASSIFICATION OF SUBJECT MATTER IPC 7 C07K14/545 C12N15/52 C12N15/63 C12N5/10 C12N15/74 C12N15/79 C07K16/24 C07K16/46 A61K38/19 C12N5/16 C12N15/62 C12Q1/68 G01N33/53 A61K9/50 A01K67/027

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

C12N A61K IPC 7

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

EPO-Internal, WPI Data, PAJ, STRAND

C. DOCUMI	ENTS CONSIDERED 1	O BE RELEVANT

Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	EP 0 879 889 A (SMITHKLINE BEECHAM CORP) 25 November 1998 (1998-11-25)	1-20,23, 24, 33-39, 42-44, 47,48, 50-52
	the whole document	
Α	EP 0 343 684 A (SYNERGEN INC) 29 November 1989 (1989-11-29) abstract page 4, line 1-36; examples 1-10 & US 5 075 222 A 24 December 1991 (1991-12-24) cited in the application	1-17
	-/	

Further documents are listed in the continuation of box C.	Patent family members are listed in annex.
Special categories of cited documents: 'A' document defining the general state of the art which is not considered to be of particular relevance 'E' earlier document but published on or after the international filing date 'L' document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) 'O' document referring to an oral disclosure, use, exhibition or other means 'P' document published prior to the international filing date but later than the priority date claimed	 "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art. "&" document member of the same patent family
Date of the actual completion of the international search 3 May 2001	Date of mailing of the international search report 18/05/2001
Name and mailing address of the ISA European Patent Office, P.B. 5818 Patentlaan 2 NL. – 2280 HV Rijswijk Tel. (+31–70) 340–2040, Tx. 31 651 epo nl, Fax: (+31–70) 340–3016	Authorized officer Mateo Rosell, A.M.

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International A tion No PCT/US 00/32400

Continuous Consideration Consideration Collegery	C (Continue	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	FC1/US 00/32400
P,X NUCLEOTIDE AND PROTEIN SEQUENCES DATABASE,			Relevant to claim No.
Clone RP11-339F22, WORKING DRAFT SEQUENCE, 5 unordered pieces. HTG; HTGS_DRAFT; HTGS_PHASEI. (from nt 76663-77367).	Ρ,Χ	EBI, UK. 14-12-99. AC = ACO16724. Homo sapiens chromosome 2	
the interleukin-1 superfamily"		<pre>clone RP11-339F22, WORKING DRAFT SEQUENCE, 5 unordered pieces. HTG; HTGS_DRAFT; HTGS_PHASE1. (from nt 76663-77367). XP002166437</pre>	
INITIAL CHARACTERIZATION OF FOUR NOVEL MEMBERS OF INTERLEUKIN-1 FAMILY" JOURNAL OF BIOLOGICAL CHEMISTRY, AMERICAN SOCIETY OF BIOLOGICAL CHEMISTS, BALTIMORE, MD, US, vol. 275, no. 14, 7 April 2000 (2000-04-07), pages 10308-10314, XP000938731 ISSN: 0021-9258 the whole document WO 00 71720 A (SIMS JOHN E ;IMMUNEX CORP (US); RENSHAW BLAIR R (US)) 34, 36-39, 42-44, 47,48, 50-52 SEQ.ID.N.1	P,X	the interleukin-1 superfamily" JOURNAL OF BIOLOGICAL CHEMISTRY, AMERICAN SOCIETY OF BIOLOGICAL CHEMISTS, BALTIMORE, MD,US, vol. 275, no. 2, 14 January 2000 (2000-01-14), pages 1169-1175, XP002135001 ISSN: 0021-9258	37-39, 42-44, 47,48,
(US); RENSHAW BLAIR R (US)) 30 November 2000 (2000-11-30) 24,33, 34, 36-39, 42-44, 47,48, 50-52	P,X	INITIAL CHARACTERIZATION OF FOUR NOVEL MEMBERS OF INTERLEUKIN-1 FAMILY" JOURNAL OF BIOLOGICAL CHEMISTRY, AMERICAN SOCIETY OF BIOLOGICAL CHEMISTS, BALTIMORE, MD, US, vol. 275, no. 14, 7 April 2000 (2000-04-07), pages 10308-10314, XP000938731 ISSN: 0021-9258	34, 36-39, 42-44, 47,48,
SEQ.ID.N.1	E	(US); RENSHAW BLAIR R (US))	24,33, 34, 36-39, 42-44, 47,48,
			30 32
		•	

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FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Continuation of Box I.1

Although claims 33,47 and 52 are directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.

Although claim(s) 48 are directed to a diagnostic method practised on the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.

Internation plication No. PCT/US 00/32400

Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. X Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
see FURTHER INFORMATION sheet PCT/ISA/210
2. Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
Dox ii Observations where unity of invention is facking (Continuation of item 2 of inst sheet)
This International Searching Authority found multiple inventions in this international application, as follows:
As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
Remark on Protest The additional search fees were accompanied by the applicant's protest.
No protest accompanied the payment of additional search fees.

Information on patent family members

International A ation No
PCT/US 00/32400

Patent documen cited in search rep		Publication date	Patent family member(s)	Publication date
EP 0879889	Α	25-11-1998	US 5945310 A CA 2221866 A JP 11000177 A JP 2000083688 A	31-08-1999 19-11-1998 06-01-1999 28-03-2000
EP 0343684	A	29-11-1989	AT 88502 T AU 633831 B AU 3746989 A BR 8907457 A DD 283835 A DE 68906077 D DE 68906077 T DE 343684 T DK 8591 A EP 0541920 A ES 2018750 T GR 90300115 T IE 66712 B IE 950317 L JP 3505279 T JP 2001029093 A KR 148009 B KR 9702917 B MX 16206 A NO 905090 A NZ 229326 A OA 9631 A SG 59969 A WO 8911540 A US 5075222 A ZA 8904052 A	15-05-1993 11-02-1993 12-12-1989 02-04-1991 24-10-1990 27-05-1993 29-07-1993 08-11-1990 18-01-1991 19-05-1993 16-10-1994 27-09-1991 24-01-1996 27-11-1989 21-11-1991 06-02-2001 01-08-1998 12-03-1997 01-11-1993 28-01-1991 23-12-1992 30-04-1993 22-02-1999 30-11-1989 24-12-1991 25-04-1990